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! Ado44524 Human GLP-1 peptide derivative 8S-
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1 Ado44523 Human GLP-1 peptide derivative 8S-
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ! Ado44526 Human GLP-1 peptide derivative 8S
                                                                                                                                                                                                                                               H(A,S)BGTFTSDVSSYLEGQAA(K,Q)BFIAWLV(N,K)G(X){0,}(R,K)(K){1,9}R
H(S)BGTFTSDVSSYLEGQAA(K)BFIAWLV(K)G(K)K{4}R
HSEGTFTSDVSSYLEGQAAKEFIAWLVKGKKKKKR
                                                                                                                                                                                                                                                                                                                                                                                                                                                                  H(A,S)EGTFTSDVSSYLEGGAA(K,Q)EPIAWLV(N,K)G(X){0,}(R,K)(K){1,9}R
H(S)EGTFTSDVSSYLEGGAA(K)EPIAWLVKGKKKKKKK
HSEGTFTSDVSSYLEGGAAKEPIAWLVKGKKKKKKKR
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     H(A, S) EGTFTSDVSSYLEGQAA(K, Q) BFIAMLV(N, K) G(X) {0,} {R, K) (K) {1,9}R
H(S) EGTFTSDVSSYLEGQAA(K) BFIAMLV(K) G(K) K{2}R
HSEGTFTSDVSSYLEGQAAKEFIAMLVKGKKKR
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        H(A,S)EGTFTSDVSSYLEGQAA(K,Q)EFIAMLV(N,K)G(X){0,}(R,K)(K){1,9}R
H(S)EGTFTSDVSSYLEGQAA(Q)EPIAMLV(N)G(K)K{4}R
HSEGTFTSDVSSYLEGQAAQEFIAMLVNGKKKGGR
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          H(A,S)EGTFTSDVSSYLEGQAA(K,Q)EFIAWLV(N,K)G(X){0,}{R,K)(K){1,9}R
H(S)EGTFTSDVSSYLEGQAA(K)EFIAWLV(K)G(K)KR
HSEGTFTSDVSSYLEGQAAKEFIAWLVKGKKR
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             H(A,S)EGTFTSDVSSYLEGQAA(K,Q)EFIAMLV(N,K)G(X){0,}{R,K).(K){1,9}R
H(S)EGTFTSDVSSYLEGQAA(K)EFIAMLV(K)G(K)K{9}R
HSEGTFTSDVSSYLEGQAAKEFIAMLVKGKKKKKKKKKRR
                                                         1 (R, K) (K) {1,9}R(X) {0,}H(A,S) EGTFTSDVSSYLEGQAA(K,Q) BFIAWLV(N,K)G 2 H(A,S) EGTFTSDVSSYLEGQAA(K,Q) BFIAWLV(N,K)G(X) {0,} (R,K) (K) {1,9}R
| FINDPATTERNS on geneseqp: * allowing 0 mismatches
                                                                                                                                                                                         ck: 7606 len: 35
                                                                                                                                                                                                                                                                                                                                                                                                            ck: 3095 len: 37
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ck: 2417 len: 33
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                                                                                                                                                                                         AD044524
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EMBL, Release 8.0, Released on 4Apr2006, Formatted on 29Apr2006

Databases searched:

6 457,216,429 2,589,679 10:15.32

Total finds:
Total length:
Total sequences:
CPU time:

```
New glucagon-like peptide 1 derivatives comprising an added C-terminal peptide, with improved transmucosal absorbability used for the treatment of diabetes.
                                                                                        GLP-1, glucagon-like peptide 1; dipeptidylpeptidase IV; trypsin;
antidiabetic; anorectic; insulin secretion.
                                                                     Human GLP-1 peptide derivative 8S-des36R-GLP1+2KR.
                                                                                                                                                                       /note= "C-terminal amide"
                                                                                                                                                                                                                                                                         (SANW ) SANWA KAGAKU KENKYUSHO CO LID.
                                                                                                                                                    Location/Qualifiers
11AA_SEQUENCE 1.0
ID ADO44522 standard; peptide; 32 AA.
                                                                                                                                                                                                                                                                                              Kouzaki T,
                                                                                                                                                                                                                                  10-OCT-2003; 2003WO-JP013020.
                                                                                                                                                                                                                                                      11-OCT-2002; 2002JP-00299283.
                                                 29-JUL-2004 (first entry)
                                                                                                                                                                                                                                                                                             Hayashi Y, Makino M,
                                                                                                                                                                                                                                                                                                                WPI; 2004-357426/33.
                                                                                                                                                                                           WO2004037859-A1
                                                                                                                                                             Modified-site
                                                                                                                      Homo sapiens
Synthetic.
                                                                                                                                                                                                               36-MAY-2004
                              ADO44522;
                               BXBXBXXXXXXBXBXBXB
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Example 1; SEQ ID NO 13; 48pp; Japanese

Takeda M, Jomori T;

Yaa, where Waa is arginine or lysine; Xaa is arginine or lysine; Yaa is arginine, arginine amide, lysine, lysine amide or homoserine; and n is 0-14. The GLP-1 peptide derivatives have to locarace to dipeptidylpeptidase IV and to trypsin due to the nature of the substitution. The peptides can be synthesised by standard solid-state peptide synthesis methods. The epptides can be used in the treatment of diabetes (insulin-dependent or insulin non-dependent), obesity and excessive appetite. Sequences ADO44512-ADO44534 represent examples of GLP-1 peptide derivatives. The invention relates to peptides consisting of a sequence derived from alucagon-like peptide 1 (GLP-1) residues 7.35 by addition, deletion and/or substitution of one or more amino acid residues. The GLP-1 derived peptides have an added sequence at the C-terminal of formula Waa (Xaal)n-

Sequence 32 AA;

ADO44522 Length: 32 June 8, 2007 16:30 Type: P Check: 9935

1 HSEGTFTSDV SSYLEGQAAK BFIAWLVKGK KR

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Yaa, where Waa is arginine or lysine; Xaa is arginine or lysine; Yaa is arginine, arginine amide, lysine, lysine amide or homoserine; and n is 0-14. The GLP-1 peptide derivatives have tolerance to dipeptidylpeptidase IV and to trypsin due to the nature of the substitution. The peptides can be synthesised by standard solid-state peptide synthesis methods. The peptides can be used in the treatment of diabetes (insulin-dependent or insulin non-dependent), obesity and excessive appetite. Sequences ADO44512-ADO44534 represent examples of GLP-1 peptide derivatives.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          The invention relates to peptides consisting of a sequence derived from glucagon-like peptide I (GLP-1) residues 7.35 by addition, deletion and/or substitution of one or more amino acid residues. The GLP-1 derived peptides have an added sequence at the C-terminal of formula Waa-(Xaa)n-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         New glucagon-like peptide 1 derivatives comprising an added C-terminal peptide, with improved transmucosal absorbability used for the treatment of diabetes.
                                                                                                                                                                                                            GLP-1; glucagon-like peptide 1; dipeptidylpeptidase IV; trypsin;
antidiabetic; anorectic; insulin secretion.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Kouzaki T, Takeda M, Jomori T;
                                                                                                                                                               Human GLP-1 peptide derivative 8S-des36R-GLP1+3XR
                                                                                                                                                                                                                                                                                                                                                                                                 /note= "C-terminal amide"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 1; SEQ ID NO 14; 48pp; Japanese
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (SANW ) SANWA KAGAKU KENKYUSHO CO LID.
                                                                                                                                                                                                                                                                                                                                     Location/Qualifiers
33
!!AA_SEQUENCE 1.0
ID AD044523 standard; peptide; 33 AA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   10-OCT-2003; 2003WO-JP013020.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   11-OCT-2002; 2002JP-00299283
                                                                                                                  29-JUL-2004 (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               eptides have an added
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Hayashi Y, Makino M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        WPI; 2004-357426/33.
                                                                                                                                                                                                                                                                                                                                                                                                                                           WO2004037859-A1
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 33 AA
                                                                                                                                                                                                                                                                                                                                                   Key
Modified-site
                                                                                                                                                                                                                                                                              Homo sapiens
Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        06-MAY-2004
                                                                    ADO44523;
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ADO44523 Length: 33 June 8, 2007 16:31 Type: P Check: 2417

1 HSEGTFTSDV SSYLEGQAAK EFIAWLVKGK KKR

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11AA_SEQUENCE 1.0
ID ADO44524 standard; peptide; 35 AA.
                                                                                            29-JUL-2004 (first entry
                                                          ADO44524;
```

GLP-1; glucagon-like peptide 1; dipeptidylpeptidase IV; trypsin; antidiabetic; anorectic; insulin secretion. Human GLP-1 peptide derivative 8S-des36R-GLP1+5KR.

Homo sapiens. Synthetic.

. Location/Qualifiers Key Modified-site

/note= "C-terminal amide"

WO2004037859-A1

06-MAY-2004

10-OCT-2003; 2003WO-JP013020.

11-OCT-2002; 2002JP-00299283

Jomori T; Hayashi Y, Makino M, Kouzaki T, Takeda M,

(SANW) SANWA KAGAKU KENKYUSHO CO LID.

New glucagon-like peptide I derivatives comprising an added C-terminal peptide, with improved transmucosal absorbability used for the treatment of diabetes. WPI; 2004-357426/33.

Example 1; SEQ ID NO 15; 48pp; Japanese.

The invention relates to peptides consisting of a sequence derived from glucagon-like peptide 1 (GLP-1) residues 7-35 by addition, deletion and/or substitution of one or more amino acid residues. The GLP-1 derived peptides have an added sequence at the C-terminal of formula Waa-(Xaa)n-Yaa, where Waa is arginine or lysine, Xaa is arginine, arginine amide, lysine, lysine amide or homoserine; and n is 0-14. The GLP-1 peptide derivatives have tolerance to dipeptidylpeptidase IV and to trypsin due to the nature of the substitution. The peptides can be synthesised by standard solid-state peptide synthesis methods. The peptides can be used in the treatment of diabetes (insulin-dependent or insulin non-dependent), obesity and excessive appetite. Sequences

Sequence 35 AA;

ADO44524 Length: 35 June 8, 2007 16:31 Type: P Check: 7606

1 HSEGTFTSDV SSYLEGQAAK EFIAWLVKGK KKKKR

```
!!AA_SEQUENCE 1.0
ID ADO44525 standard; peptide; 37 AA.
                                                      ADO44525;
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29-JUL-2004 (first entry)

Human GLP-1 peptide derivative 8S-des36R-GLP1+7KR.

GLP-1; glucagon-like peptide 1; dipeptidylpeptidase IV; trypsin; antidiabėtic; anorectic; insulin secretion.

Homo sapiens. Synthetic.

/note= "C-terminal amide" Location/Qualifiers Modified-site

WO2004037859-A1

06-MAY-2004

10-OCT-2003; 2003WO-JP013020.

11-OCT-2002; 2002JP-00299283

(SANW) SANWA KAGAKU KENKYUSHO CO LTD.

Jomori T; Kouzaki T, Takeda M, Hayashi Y, Makino M,

WPI; 2004-357426/33

New glucagon-like peptide I derivatives comprising an added C-terminal peptide, with improved transmucosal absorbability used for the treatment of diabetes.

Example 1; SEQ ID NO 16; 48pp; Japanese

Yaa, where Waa is arginine or lysine; Xaa is arginine or lysine; Yaa is arginine, arginine amide, lysine, lysine amide or homoserine; and n is 0-14. The GLP-1 peptide derivatives have tolerance to dipeptidylpeptidase be synthesised by standard solid-state peptide synthesis methods. The peptides can peptides can be used in the treatment of diabetes (insulin-dependent or insulin non-dependent), obesity and excessive appetite. Sequences ADO44512-ADO44534 represent examples of GLP-1 peptide derivatives. The invention relates to peptides consisting of a sequence derived from dilucagon-like peptide 1 (GLP-1) residues 7-35 by addition, deletion and/or substitution of one or more amino acid residues. The GLP-1 derived peptides have an added sequence at the C-terminal of formula Waa (Xaa)n-

Sequence 37 AA;

ADO44525 Length: 37 June 8, 2007 16:31 Type: P Check: 3095

1 HSEGTFTSDV SSYLEGOAAK EFIAWLVKGK KKKKKKR

ado44526.geneseqp20048

```
Human GLP-1 peptide derivative 8S-des36R-GLP1+10XR.
!!AA_SEQUENCE 1.0
ID ADO44526 standard; peptide; 40 AA.
                                                                                                                             29-JUL-2004 (first entry)
                                                                           ADO44526;
```

GLP-1; glucagon-like peptide 1; dipeptidylpeptidase IV; trypsin; antidiabetic; anorectic; insulin secretion. Homo sapiens. Synthetic.

/note= "C-terminal amide" Location/Qualifiers Modified-site

WO2004037859-A1

10-OCT-2003; 2003WO-JP013020.

06-MAY-2004

11-OCT-2002; 2002JP-00299283.

SANW) SANWA KAGAKU KENKYUSHO CO LTD.

Kouzaki T, Takeda M, Jomori T; Makino M, Hayashi Y,

WPI; 2004-357426/33.

New glucagon-like peptide I derivatives comprising an added C-terminal peptide, with improved transmucosal absorbability used for the treatment of diabetes.

Example 1; SEQ ID NO 17; 48pp; Japanese.

Yaa, where Waa is arginine or lysine; Xaa is arginine or lysine; Yaa is arginine, arginine amide, lysine, lysine amide or homoserine; and n is 0-14. The GLP-1 peptide derivatives have tolerance to dipeptidylpeptidase IV and to trypsin due to the nature of the substitution. The peptides can be synthesised by standard solid-state peptide synthesis methods. The peptides can be used in the treatment of diabetes (insulin-dependent or insulin non-dependent), obesity and excessive appetite. Sequences ADO44512-ADO44534 represent examples of GLP-1 peptide derivatives. The invention relates to peptides consisting of a sequence derived from glucagon-like peptide 1 (GLP-1) residues 7-35 by addition, deletion and/or substitution of one or more amino acid residues. The GLP-1 derived peptides have an added sequence at the C-terminal of formula Waa-(Xaaln-

Sequence 40 AA;

ADO44526 Length: 40 June 8, 2007 16:30 Type: P Check: 1891

1 HSEGTFTSDV SSYLEGQAAK EFIAWLVKGK KKKKKKKKK

```
IIAA_SEQUENCE 1.0
ID ADO44532 standard; peptide; 35 AA.
```

AD044532;

29-JUL-2004 (first entry)

Human GLP-1 peptide derivative 8S26Q34N-des36R-GLP1-5KR.

GLP-1; glucagon-like peptide 1; dipeptidylpeptidase IV; trypsin; antidiabetic; anorectic; insulin secretion.

Homo sapiens Synthetic.

Location/Qualifiers 35 Key Modified-site

/note= "C-terminal amide"

WO2004037859-A1

06-MAY-2004

10-OCT-2003; 2003WO-JP013020.

11-OCT-2002; 2002JP-00299283

(SANW) SANWA KAGAKU KENKYUSHO CO LTD.

Kouzaki T, Takeda M, Jomori T; Hayashi Y, Makino M,

WPI; 2004-357426/33.

New glucagon-like peptide 1 derivatives comprising an added C-terminal peptide, with improved transmucosal absorbability used for the treatment of diabetes.

Example 1; SEQ ID NO 23; 48pp; Japanese

The invention relates to peptides consisting of a sequence derived from glucagon-like peptide 1 (GLP-1) residues 7-35 by addition, deletion and/or substitution of one or more amino acid residues. The GLP-1 derived peptides have an added sequence at the C-terminal of formula Waa-(Xaa)n-Yaa, where Waa is arginine or lysine; Xaa is arginine, arginine amide, lysine, lysine amide or homoserine; and n is 0-14. The GLP-1 peptide derivatives have tolerance to dipeptidylpeptidase IV and to trypsin due to the nature of the substitution. The peptides can be synthesised by standard solid-state peptide synthesis methods. The peptides can be used in the treatment of diabetes (insulin-dependent or peptides can be used in the treatment of diabetes (insulin-dependinsulin non-dependent), obesity and excessive appetite. Sequences ADO44512-ADO44534 represent examples of GLP-1 peptide derivatives

Sequence 35 AA;

ADO44532 Length: 35 June 8, 2007 16:31 Type: P Check: 7810

1 HSEGTFTSDV SSYLEGQAAQ EFIAWLVNGK KKKKK

! FINDPATTERNS on uniprot: * allowing 0 mismatches

1 (R, K) (K) {1,9}R(X) {0,}H(A,S) EGTFTSDVSSYLECQAA(K,Q) EFIAWLV (N, K) G 2 H(A,S) EGTFTSDVSSYLEGQAA(K,Q) EFIAWLV (N, K) G(X) {0,} (R, K) (K) {1,9}R

Databases searched: UNIPROT, Release 7.2, Released on 7Mar2006, Formatted on 7Mar2006

Total finds: Total length: Total sequences: CPU time:

925,015,592 2,849,598 17:54.40

| FINDPATTERNS on pir: * allowing 0 mismatches

1 (R, K) (K) {1,9}R(X) {0,}H(A,S) EGTFTSDVSSYLEGGAA (K,Q) EFIAWLV (N,K)G 2 H(A,S) EGTFTSDVSSYLEGGAA (K,Q) EFIAWLV (N,K)G(X) {0,} (R,K) (K) {1,9}R

Databases searched: NBRP, Release 80.0, Released on 31Dec2004, Formatted on 21Jun2005

0 96,216,763 283,416 01:50.42

Total finds: Total length: Total sequences: CPU time: